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## NOTICE OF ALLOWANCE AND FEE(S) DUE

GE HEALTHCARE BIO-SCIENCES CORP.
MELISSA LECK
101 CARNEGIE CENTER
PATENT DEPARTMENT
PRINCETON, NJ 08540

EXAMINER

MONSHIPOURI, MARYAM

ART UNIT PAPER NUMBER

1656

DATE MAILED: 06/15/2011

| APPLICATION NO. | FILING DATE | FIRST NAMED INVENTOR | ATTORNEY DOCKET NO. | CONFIRMATION NO. |
|-----------------|-------------|----------------------|---------------------|------------------|
| 10/583,591      | 06/19/2006  | Simon L. Stubbs      | PA0394              | 4011             |

TITLE OF INVENTION: CYTOCHROME C PROTEIN AND ASSAY

| APPLN. TYPE    | SMALL ENTITY | ISSUE FEE DUE | PUBLICATION FEE DUE | PREV. PAID ISSUE FEE | TOTAL FEE(S) DUE | DATE DUE   |
|----------------|--------------|---------------|---------------------|----------------------|------------------|------------|
| nonprovisional | NO           | \$1510        | \$300               | \$0                  | \$1810           | 09/15/2011 |

THE APPLICATION IDENTIFIED ABOVE HAS BEEN EXAMINED AND IS ALLOWED FOR ISSUANCE AS A PATENT. PROSECUTION ON THE MERITS IS CLOSED. THIS NOTICE OF ALLOWANCE IS NOT A GRANT OF PATENT RIGHTS. THIS APPLICATION IS SUBJECT TO WITHDRAWAL FROM ISSUE AT THE INITIATIVE OF THE OFFICE OR UPON PETITION BY THE APPLICANT. SEE 37 CFR 1.313 AND MPEP 1308.

THE ISSUE FEE AND PUBLICATION FEE (IF REQUIRED) MUST BE PAID WITHIN <u>THREE MONTHS</u> FROM THE MAILING DATE OF THIS NOTICE OR THIS APPLICATION SHALL BE REGARDED AS ABANDONED. <u>THIS STATUTORY PERIOD CANNOT BE EXTENDED.</u> SEE 35 U.S.C. 151. THE ISSUE FEE DUE INDICATED ABOVE DOES NOT REFLECT A CREDIT FOR ANY PREVIOUSLY PAID ISSUE FEE IN THIS APPLICATION. IF AN ISSUE FEE HAS PREVIOUSLY BEEN PAID IN THIS APPLICATION (AS SHOWN ABOVE), THE RETURN OF PART B OF THIS FORM WILL BE CONSIDERED A REQUEST TO REAPPLY THE PREVIOUSLY PAID ISSUE FEE TOWARD THE ISSUE FEE NOW DUE.

#### HOW TO REPLY TO THIS NOTICE:

I. Review the SMALL ENTITY status shown above.

If the SMALL ENTITY is shown as YES, verify your current SMALL ENTITY status:

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B. If the status above is to be removed, check box 5b on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and twice the amount of the ISSUE FEE shown above, or

If the SMALL ENTITY is shown as NO:

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B. If applicant claimed SMALL ENTITY status before, or is now claiming SMALL ENTITY status, check box 5a on Part B - Fee(s) Transmittal and pay the PUBLICATION FEE (if required) and 1/2 the ISSUE FEE shown above.

II. PART B - FEE(S) TRANSMITTAL, or its equivalent, must be completed and returned to the United States Patent and Trademark Office (USPTO) with your ISSUE FEE and PUBLICATION FEE (if required). If you are charging the fee(s) to your deposit account, section "4b" of Part B - Fee(s) Transmittal should be completed and an extra copy of the form should be submitted. If an equivalent of Part B is filed, a request to reapply a previously paid issue fee must be clearly made, and delays in processing may occur due to the difficulty in recognizing the paper as an equivalent of Part B.

III. All communications regarding this application must give the application number. Please direct all communications prior to issuance to Mail Stop ISSUE FEE unless advised to the contrary.

IMPORTANT REMINDER: Utility patents issuing on applications filed on or after Dec. 12, 1980 may require payment of maintenance fees. It is patentee's responsibility to ensure timely payment of maintenance fees when due.

### PART B - FEE(S) TRANSMITTAL

#### Complete and send this form, together with applicable fee(s), to: Mail Mail Stop ISSUE FEE

Commissioner for Patents
P.O. Box 1450
Alexandria, Virginia 22313-1450
or Fax (571)-273-2885

| naintenance fee notificat   | tions.  | or transmitting the 1880 ag the Patent, advance of the transmitting the 1880 are transmitting the 1880 and the 1880 are transmitting the 1880 are transmitted to 1880 are transmitted the 1880 a |  |   |  |  |  |   |  |
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| CURRENT CORRESPONDI  22840  | 7590 06/15  | ock 1 for any change of address)   | F  | ∃ee(s]<br>oaper   | ) Transmittal. This<br>s. Each additional  | certif                                       | g can only be used for<br>icate cannot be used for<br>, such as an assignment<br>ling or transmission.                       | or any oth  | her accompanying   |
|   | CARE BIO-SCIEN<br>K<br>E CENTER   |  | I<br>S<br>a<br>u   | here<br>States<br>iddre<br>ransr  | Certi<br>by certify that this<br>Postal Service wi<br>ssed to the Mail<br>mitted to the USPT             | ificate<br>Fee(s<br>th suf<br>Stop<br>O (57  | of Mailing or Transı<br>s) Transmittal is being<br>ficient postage for firs<br>ISSUE FEE address<br>1) 273-2885, on the da   | nission<br>deposite<br>t class ma<br>above, o<br>te indicat | d with the United<br>ail in an envelope<br>r being facsimile<br>and below.                           |
| PRINCETON, N  | IJ 08540  |  |  |   |  |  |  |   | (Depositor's name)   |
|   |   |  | Ļ  |   |  |  |  |   | (Signature)  |
|   |   |  | L  |   |  |  |  |   | (Date)   |
| APPLICATION NO.   | FILING DATE   |  | FIRST NAMED INVENT   | OR  |  | ATTO:  | RNEY DOCKET NO.  | CONFI   | RMATION NO.  |
| 10/583,591  | 06/19/2006  |  | Simon L. Stubbs  |   |  |  | PA0394   |   | 4011   |
| ITLE OF INVENTION   | : CYTOCHROME C PR   |  |  |   |  |  |  |   |  |
| APPLN. TYPE   | SMALL ENTITY  | ISSUE FEE DUE  | PUBLICATION FEE DU   | Æ   | PREV. PAID ISSUE   | FEE  | TOTAL FEE(S) DUE   |   | DATE DUE   |
| nonprovisional  | NO  | \$1510   | \$300  | 00 \$0  |  | \$1810                                       |  | (   | 09/15/2011   |
| EXAM  | EXAMINER ART UNIT CLASS-SUBCI   |  |  |   |  |  |  |   |  |
| MONSHIPOUI  | RI, MARYAM  | 1656   | 436-086000   |   |  |  |  |   |  |
| Change of correspondence address or indication of "Fee Address" (37 FR 1.363).  Change of correspondence address (or Change of Correspondence Address form PTO/SB/122) attached.  "Fee Address" indication (or "Fee Address" Indication form PTO/SB/47; Rev 03-02 or more recent) attached. Use of a Customer Number is required. |   |  | (1) the names of up<br>or agents OR, altern<br>(2) the name of a si<br>registered attorney of<br>2 registered patent a     | 2. For printing on the patent front page, list  (1) the names of up to 3 registered patent attorneys or agents OR, alternatively,  (2) the name of a single firm (having as a member a registered attorney or agent) and the names of up to 2 registered patent attorneys or agents. If no name is listed, no name will be printed. |  |  |  |   |  |
| PLEASE NOTE: Unl<br>recordation as set forth<br>(A) NAME OF ASSIG   | ess an assignee is ident<br>h in 37 CFR 3.11. Comp<br>GNEE  | A TO BE PRINTED ON The strength of this form is NO strength of this form is NO strength of the | data will appear on the T a substitute for filing (B) RESIDENCE: (CI   | e pat<br>an as  | ent. If an assignerssignment. and STATE OR CO  | DUNT   |  |   | _  |
|   | are submitted:  fo small entity discount profes   | permitted)   | o. Payment of Fee(s): (F   | d.<br>card  | . Form PTO-2038 i  | is atta                                      | ched.  |   | ,  |
| a. Applicant claims   | t <b>us</b> (from status indicated<br>s SMALL ENTITY statu  | is. See 37 CFR 1.27.   | ☐ b. Applicant is no   | longe   | er claiming SMAL   | L ENT  | FITY status. See 37 CF   | R 1.27(g  | g)(2).   |
| OTE: The Issue Fee and<br>terest as shown by the r  | d Publication Fee (if requeecords of the United Sta   | uired) will not be accepte<br>tes Patent and Trademark   | d from anyone other that<br>Office.  | an the  | e applicant; a regis   | tered a                                      | attorney or agent; or th   | e assigne   | e or other party in  |
| Authorized Signature  |   |  |  |   | Date   |  |  |   |  |
| Typed or printed name   |   |  |  |   | _  |  |  |   |  |
| nis collection of inform<br>application. Confident<br>abmitting the completed<br>is form and/or suggesti<br>ox 1450, Alexandria, V<br>lexandria, Virginia 223   | ation is required by 37 C<br>tiality is governed by 35<br>I application form to the<br>ons for reducing this bur<br>irginia 22313-1450. DO<br>13-1450 | FR 1.311. The informatic<br>U.S.C. 122 and 37 CFR<br>USPTO. Time will vary<br>rden, should be sent to th<br>NOT SEND FEES OR (   | on is required to obtain<br>1.14. This collection is<br>depending upon the in<br>e Chief Information Of<br>COMPLETED FORMS | or re<br>estir<br>divid<br>ficer<br>TO  | tain a benefit by th<br>mated to take 12 m<br>dual case. Any con<br>, U.S. Patent and T<br>THIS ADDRESS. | e publ<br>iinutes<br>nment<br>'raden<br>SENI | ic which is to file (and<br>to complete, includin<br>s on the amount of tin<br>ark Office, U.S. Depa<br>D TO: Commissioner f | by the Ug gathering you re you re or Patent                 | SPTO to process)<br>ng, preparing, and<br>quire to complete<br>f Commerce, P.O.<br>s, P.O. Box 1450, |

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| APPLICATION NO.                | FILING DATE     | FIRST NAMED INVENTOR | ATTORNEY DOCKET NO. | CONFIRMATION NO. |  |
|--------------------------------|-----------------|----------------------|---------------------|------------------|--|
| 10/583,591                     | 06/19/2006      | Simon L. Stubbs      | PA0394              | 4011             |  |
| 22840 75                       | 90 06/15/2011   |                      | EXAM                | IINER            |  |
|                                | RE BIO-SCIENCES | MONSHIPOURI, MARYAM  |                     |                  |  |
| MELISSA LECK<br>101 CARNEGIE C | ENTER           | ART UNIT             | PAPER NUMBER        |                  |  |
| PATENT DEPART<br>PRINCETON, NJ |                 | 1656                 |                     |                  |  |

## Determination of Patent Term Adjustment under 35 U.S.C. 154 (b)

(application filed on or after May 29, 2000)

The Patent Term Adjustment to date is 483 day(s). If the issue fee is paid on the date that is three months after the mailing date of this notice and the patent issues on the Tuesday before the date that is 28 weeks (six and a half months) after the mailing date of this notice, the Patent Term Adjustment will be 483 day(s).

If a Continued Prosecution Application (CPA) was filed in the above-identified application, the filing date that determines Patent Term Adjustment is the filing date of the most recent CPA.

Applicant will be able to obtain more detailed information by accessing the Patent Application Information Retrieval (PAIR) WEB site (http://pair.uspto.gov).

Any questions regarding the Patent Term Extension or Adjustment determination should be directed to the Office of Patent Legal Administration at (571)-272-7702. Questions relating to issue and publication fee payments should be directed to the Customer Service Center of the Office of Patent Publication at 1-(888)-786-0101 or (571)-272-4200.

### **Privacy Act Statement**

The Privacy Act of 1974 (P.L. 93-579) requires that you be given certain information in connection with your submission of the attached form related to a patent application or patent. Accordingly, pursuant to the requirements of the Act, please be advised that: (1) the general authority for the collection of this information is 35 U.S.C. 2(b)(2); (2) furnishing of the information solicited is voluntary; and (3) the principal purpose for which the information is used by the U.S. Patent and Trademark Office is to process and/or examine your submission related to a patent application or patent. If you do not furnish the requested information, the U.S. Patent and Trademark Office may not be able to process and/or examine your submission, which may result in termination of proceedings or abandonment of the application or expiration of the patent.

The information provided by you in this form will be subject to the following routine uses:

- 1. The information on this form will be treated confidentially to the extent allowed under the Freedom of Information Act (5 U.S.C. 552) and the Privacy Act (5 U.S.C 552a). Records from this system of records may be disclosed to the Department of Justice to determine whether disclosure of these records is required by the Freedom of Information Act.
- 2. A record from this system of records may be disclosed, as a routine use, in the course of presenting evidence to a court, magistrate, or administrative tribunal, including disclosures to opposing counsel in the course of settlement negotiations.
- 3. A record in this system of records may be disclosed, as a routine use, to a Member of Congress submitting a request involving an individual, to whom the record pertains, when the individual has requested assistance from the Member with respect to the subject matter of the record.
- 4. A record in this system of records may be disclosed, as a routine use, to a contractor of the Agency having need for the information in order to perform a contract. Recipients of information shall be required to comply with the requirements of the Privacy Act of 1974, as amended, pursuant to 5 U.S.C. 552a(m).
- 5. A record related to an International Application filed under the Patent Cooperation Treaty in this system of records may be disclosed, as a routine use, to the International Bureau of the World Intellectual Property Organization, pursuant to the Patent Cooperation Treaty.
- 6. A record in this system of records may be disclosed, as a routine use, to another federal agency for purposes of National Security review (35 U.S.C. 181) and for review pursuant to the Atomic Energy Act (42 U.S.C. 218(c)).
- 7. A record from this system of records may be disclosed, as a routine use, to the Administrator, General Services, or his/her designee, during an inspection of records conducted by GSA as part of that agency's responsibility to recommend improvements in records management practices and programs, under authority of 44 U.S.C. 2904 and 2906. Such disclosure shall be made in accordance with the GSA regulations governing inspection of records for this purpose, and any other relevant (i.e., GSA or Commerce) directive. Such disclosure shall not be used to make determinations about individuals.
- 8. A record from this system of records may be disclosed, as a routine use, to the public after either publication of the application pursuant to 35 U.S.C. 122(b) or issuance of a patent pursuant to 35 U.S.C. 151. Further, a record may be disclosed, subject to the limitations of 37 CFR 1.14, as a routine use, to the public if the record was filed in an application which became abandoned or in which the proceedings were terminated and which application is referenced by either a published application, an application open to public inspection or an issued patent.
- 9. A record from this system of records may be disclosed, as a routine use, to a Federal, State, or local law enforcement agency, if the USPTO becomes aware of a violation or potential violation of law or regulation.

|   | Application No.  | Applicant(s)  |
|---|--|---|
|   | 10/583.591   | STUBBS ET AL.   |
| Notice of Allowability  | Examiner   | Art Unit  |
|   | MARYAM MONSHIPOURI   | 1656  |
| The MAILING DATE of this communication appea.  All claims being allowable, PROSECUTION ON THE MERITS IS herewith (or previously mailed), a Notice of Allowance (PTOL-85) NOTICE OF ALLOWABILITY IS NOT A GRANT OF PATENT RI of the Office or upon petition by the applicant. See 37 CFR 1.313  1. This communication is responsive to 2/10/11.  2. The allowed claim(s) is/are 1-4,12-15,21-38,46 and 47.  3. Acknowledgment is made of a claim for foreign priority under the series of th | (OR REMAINS) CLOSED in this a or other appropriate communicati IGHTS. This application is subject and MPEP 1308. | application. If not included on will be mailed in due course. <b>THIS</b> |
| a) ☑ All b) ☐ Some* c) ☐ None of the:   |  |   |
| 1. Certified copies of the priority documents have  |  |   |
| 2. Certified copies of the priority documents have  | • •  |   |
| 3. Copies of the certified copies of the priority do  | cuments have been received in thi  | s national stage application from the                                     |
| International Bureau (PCT Rule 17.2(a)).  * Certified copies not received:  |  |   |
| Applicant has THREE MONTHS FROM THE "MAILING DATE" noted below. Failure to timely comply will result in ABANDONN THIS THREE-MONTH PERIOD IS NOT EXTENDABLE.  4. A SUBSTITUTE OATH OR DECLARATION must be subminformal patent APPLICATION (PTO-152) which give   | IENT of this application. itted. Note the attached EXAMINE   | R'S AMENDMENT or NOTICE OF  |
| <ul> <li>5. ☐ CORRECTED DRAWINGS (as "replacement sheets") mus</li> <li>(a) ☐ including changes required by the Notice of Draftspers</li> </ul>   | st be submitted.   |   |
| (a) ☐ including changes required by the Notice of Dratispers  1) ☐ hereto or 2) ☐ to Paper No./Mail Date  | •  | J-946) attached   |
| (b) ☐ including changes required by the attached Examiner's Paper No./Mail Date  Identifying indicia such as the application number (see 37 CFR 1.  | s Amendment / Comment or in the  |   |
| each sheet. Replacement sheet(s) should be labeled as such in t   |  |   |
| <ol> <li>DEPOSIT OF and/or INFORMATION about the depo-<br/>attached Examiner's comment regarding REQUIREMENT</li> </ol>   |  |   |
| Attachment(s)  1. ☐ Notice of References Cited (PTO-892)  2. ☐ Notice of Draftperson's Patent Drawing Review (PTO-948)  3. ☑ Information Disclosure Statements (PTO/SB/08), Paper No./Mail Date 2/10/11   | 5. ☐ Notice of Informal 6. ☐ Interview Summa Paper No./Mail □ 7. ☐ Examiner's Amen                               | ry (PTO-413),<br>oate<br>dment/Comment                                    |
| <ol> <li>Examiner's Comment Regarding Requirement for Deposit<br/>of Biological Material</li> </ol>   | 9. Other   | nent of Reasons for Allowance   |
|   |  |   |

Serial Number: 10/583,591

Art Unit 1656

The following is an **Examiner's Statement of Reasons for Allowance**:

Claims 1-4, 12-38, 43 and 46-47 are directed to a cytochrome C-reporter fusion construct comprising:

(a) a modified cytochrome C protein derived from wild type human cytochrome C

represented by SEQ ID NO:2 and

(b) a fluorescent protein,

wherein said fusion protein targets the mitochondria and has a reduced ability to

induce apoptosis in a living cell, and

wherein said modified cytochrome C comprises the amino acid substitution

selected from the group consisting of K73A, K73L, K73R, K73G and K73X, wherein X

represents trimethylation, a nucleic acid sequence encoding said fusion construct, host

cells stably transformed with said nucleic acid a method of detecting apoptosis in a

living cell comprising the steps of:

(i) culturing a cell transformed to over-express said fusion construct and

(ii) determining the localization of said fusion construct within the cell with time;

wherein a change in localization of the fusion construct within the cell is indicative of

apoptosis.

Claimed fusion construct is free of prior art. Further the prior art fails to suggest

such specifically claimed construct. Hence, said construct is also non-obvious. Since

said fusion construct is both novel and non-obvious, a nucleic acid sequence encoding

said construct, vectors and host cells comprising said nucleic acid sequence, and a

method of use of said fusion construct as claimed are also novel and non-obvious.

Claims 1-4, 12-38, 43 and 46-47 are allowed.

2

Serial Number:10/583,591

Art Unit 1656

Any comments considered necessary by applicant must be submitted no later than the payment of the issue fee and, to avoid processing delays, should preferably accompany the issue fee. Such submissions should be clearly labeled "Comments on Statement of Reasons for Allowance."

Any inquiry concerning this communication or earlier communications from the examiner should be directed to MARYAM MONSHIPOURI whose telephone number is (571)272-0932. The examiner can normally be reached on full flex.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Rao Munjunath can be reached on (571) 272-0939. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

/Maryam Monshipouri/

Primary Examiner, Art Unit 1656

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Application/Control Number: 10/583,591 Page 2

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